(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER		see Form PCT/ISA/220
GC839-PCT	ACTION	as well	as, where applicable, Item 5 below.
International application No.	International filing date (day/mont	h/year)	(Earliest) Priority Date (day/month/year)
PCT/US2005/012270	12/04/2005		15/04/2004
Applicant			
GENENCOR INTERNATIONAL, IN	NC.		
This international search report has been according to Article 18. A copy is being tra	prepared by this International Searcansmitted to the International Burea	ching Autho u.	rity and is transmitted to the applicant
This international search report consists of	f a total of she	ets.	•
X It is also accompanied by	a copy of each prior art document	cited in this	report.
Basis of the report			
a. With regard to the language, the	· ·		
	application in the language in which	it was filed	, which is the language
of a translation of th	e international application into rnished for the purposes of internat	ional search	(Rules 12.3(a) and 23.1(b))
b. X With regard to any nucle	otide and/or amino acid sequenc	e disclosed	in the international application, see Box No. I.
2. X Certain claims were fou	nd unsearchable (See Box No. II)		
3. Unity of invention is lac	king (see Box No III)		
4. With regard to the title ,			
the text is approved as su	bmitted by the applicant		
X the text has been establis	hed by this Authority to read as foll	ows:	
ANTI-CEA SCFV-BETA-LAC	CTAMASE CONTRUCTS (CA	B MOLEC	ULES) IN ADEPT
·		•	
			:
5. With regard to the abstract,			
X the text is approved as su	ubmitted by the applicant		
the text has been establis may, within one month from	shed, according to Rule 38.2(b), by orn the date of mailing of this internation	this Authori ational sear	ty as it appears in Box No. IV. The applicant ch report, submit comments to this Authority
6. With regard to the drawings ,			
a. the figure of the drawings to be	published with the abstract is Figure	No	
as suggested by	• •		
	is Authority, because the applicant		
	is Authority, because this figure bet	ter characte	erizes the Invention
b. X none of the figures is to b	e published with the abstract		



ternational application No.

PCT/US2005/012270

Вох	No. I	Nu	ucleotide and/	or amino acid :	sequence(s) (C	ontinuation	of item 1.	b of the fire	st sheet)	
1.	With inver	regard ntion, th	to any nucleotide e international se	e and/or amino ac earch was carried	id sequence discle out on the basis o	osed in the inte of:	ernational a	pplication and	d necessary to	the claimed
	a.	type of	material							
		Х	a sequence list	ing						
			table(s) related	to the sequence	listing					
			af matarial							
	b.	x	of material							
		X	on paper							
		ِ لِــُـا	in electronic for	m			•			
	c.	time of	f filing/furnishing	•	~					
			contained in th	e international app	plication as filed					٠
			filed together w	vith the internation	al application in e	lectronic form				
		X	furnished subs	equently to this A	uthority for the pur	pose of search	h			
_	х	In o	ddition in the en	no that more than	one version or co	ny of a seguer	nce listing a	nd/or table re	Istino thereto b	nas heen filed
2.	لتنا	or fu	irnished, the req	uired statements t	that the informatio ond the applicatio	n in the subsec	quent or ad	ditional copie	s is identical to	that in the
		appi	iication as ilieu o	r does not go bey	ond the applicatio	ii ao iiica, ao a	ірргорпакс,	Were idilinori		
3.	Addi	itional c	omments:	,						
J.	Addi	illoniai ci	ommento.							
			•		•					
										•
									•	
					•					
							٠			
								•		
			•							
						•				
					•			•		·

ational application No PCT/US2005/012270

A. CLASSIFICATION OF SUBJECT MATTER
INV. C07K19/00 A61P35/00 ADD. C07K16/30

C07K14/245

A61K31/195 C12N9/86

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) C07K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, WPI Data, PAJ, EMBASE, Sequence Search

C. DOCUMENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
Y	MEYER D L ET AL: "SITE-SPECIFIC PRODRUG ACTIVATION BY ANTIBODY— -LACTAMASE CONJUGATES: PRECLINICAL INVESTIGATION OF THE EFFICACY AND TOXICITY OF DOXOXRUBICIN DELIVERED BY ANTIBODY DIRECTED CATALYSIS" BIOCONJUGATE CHEMISTRY, ACS, WASHINGTON, DC, US, vol. 6, no. 4, 1 July 1995 (1995-07-01), pages 440-446, XP000517229 ISSN: 1043-1802 page 441	1-6, 8-13, 15-20, 22-34		
Y	WO 03/055527 A (VLAAMS INTERUNIVERSITAIR INSTITUUT VOOR BIOTECHNOLOGIE VZW; REVETS, HI) 10 July 2003 (2003-07-10) example 1	1-6, 8-13, 15-20, 22-34		

X Further documents are listed in the continuation of Box C.	X See patent family annex.
* Special categories of cited documents: *A* document defining the general state of the art which is not considered to be of particular relevance *E* earlier document but published on or after the international filling date *L* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) *O* document referring to an oral disclosure, use, exhibition or other means *P* document published prior to the international filling date but later than the priority date claimed	 'T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention 'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone 'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. '8' document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
27 April 2006	18/05/2006
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,	Authorized officer Wagner, R
Fax: (+31-70) 340-3016	

2



C(Continua	tion). DOCUMENTS CONSIDERED TO BE RELEVANT	-	
Category*	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
Υ	WU A M ET AL: "Tumor localization of anti-CEA single-chain Fvs: improved targeting by non-covalent dimers" IMMUNOTECHNOLOGY, ELSEVIER SCIENCE PUBLISHERS BV, NL, vol. 2, no. 1, February 1996 (1996-02), pages 21-36, XP004052689 ISSN: 1380-2933		1-6, 8-13, 15-20, 22-34
	table 1		
Y	MCDONAGH CHARLOTTE F ET AL: "Improved yield and stability of L49-sFv-beta-lactamase, a single-chain antibody fusion protein for anticancer prodrug activation, by protein		5,6,12, 13
	engineering." BIOCONJUGATE CHEMISTRY, vol. 14, no. 5, 25 July 2003 (2003-07-25), pages 860-869, XP002378783 ISSN: 1043-1802 page 862		
A	STICKLER M M ET AL: "CD4+ T-CELL EPITOPE DETERMINATION USING UNEXPOSED HUMAN DONOR PERIPHERAL BLOOD MONONUCLEAR CELLS" JOURNAL OF IMMUNOTHERAPY, LIPPINCOTT WILLIAMS & WILKINS, HAGERSTOWN, MD, US, vol. 23, no. 6, November 2000 (2000-11), pages 654-660, XP008040520 ISSN: 1524-9557 the whole document		7,14
E	WO 2005/058236 A (GENENCOR INTERNATIONAL, INC; FOX, JUDITH, A; HARDING, FIONA, A; SCHELL) 30 June 2005 (2005-06-30) figure 25A		1-34
A	HEFTA L J F ET AL: "Kinetic and affinity constants of epitope specific anti-carcinoembryonic antigen (CEA) monoclonal antibodies for CEA and engineered CEA domain constructs" IMMUNOTECHNOLOGY, ELSEVIER SCIENCE PUBLISHERS BV, NL, vol. 4, no. 1, June 1998 (1998-06), pages 49-57, XP004127386 ISSN: 1380-2933 the whole document		1-34
	-/		
		• •	



ategory*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
	HARDING FIONA A ET AL: "A beta-lactamase with reduced immunogenicity for the targeted delivery of chemotherapeutics using antibody-directed enzyme prodrug therapy" MOLECULAR CANCER THERAPEUTICS, vol. 4, no. 11, November 2005 (2005-11), pages 1791-1800, XP002378784 ISSN: 1535-7163 the whole document		1-34
1	WO 91/01990 A (CITY OF HOPE) 21 February 1991 (1991-02-21) the whole document		1-34
			·
			·

ternational application No. PCT/US2005/012270

Box II	Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This Int	ternational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Although claims 15-34 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
з. [Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This In	ternational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remar	k on Protest The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.

ation on patent family members

national application No PCT/US2005/012270

	tent document in search report		Publication date		Patent family member(s)		Publication date
WO	03055527	A	10-07-2003	AU CA JP US	2002361236 2471645 2005517674 2005048060	A1 T	15-07-2003 10-07-2003 16-06-2005 03-03-2005
WO	2005058236	Α	30-06-2005	NONE			
WO	9101990	A .	21-02-1991	AU AU CA DE DE JP JP US	629401 6344790 2035899 69018801 69018801 0436016 2966924 4502258 5081235	A1 D1 T2 A1 B2 T	01-10-1992 11-03-1991 27-01-1991 24-05-1995 19-10-1995 10-07-1991 25-10-1999 23-04-1992 14-01-1992